H36.D2.B7 ANTI-TISSUE FACTOR LIGHT CHAIN VARIABLE REGION

GACATTCAGATGACCCAGTCTCCTGCCTCCCAGTCTGCATCTCTGGGAGAAAGTGTCACCATCACATGC D I Q M T Q S P A S Q S A S L G E S V T I T C

CTGGCAAGTCAGACCATTGATACATGGTTAGCATGGTATCAGCAGAAACCAGGGAAATCTCCTCAGCTC

CTGATTTATGCTGCCACCAACTTGGCAGATGGGGTCCCATCAAGGTTCAGTGGCAGTGGATCTGGCACA

AAATTITCTITCAAGATCAGGCTACAGGCTGAAGATTTTGTAAATTATT TACTGT<u>CAACAAGTTTAC</u> K F S F K I S S L Q A E D F V N Y Y C Q Q V Y

AGTTCTCCATTCACGTTCGGTGCTGGGACCAAGCTGGAGA FIFGAGTKLELK

FIG. 1A

H36.D2.B7 ANTI-TISSUE FACTOR HEAVY CHAIN VARIABLE REGION

GAGATCCAGCTGCAGCAGTCTGGACCTGAGCTGGAGCCTGGGGCTTCAGTGCAGGTATCCTGCAAG E I Q L Q Q S G P E L V K P G A S V Q V S C

ACTICTGGITACTCATTCACTGACTACAGGTGAGGCAGAGAGCCATGGAAAGAGCCTTGAG SGYSFIDYNVYWVRQSHGKSL

TGGATTGGA<u>TATATTGATCCTTACAATGGTATTACTATCTACGACCAGAACTTCAAGGGC</u>AAGGCCACA GYIDPYNGITIYDONFKGKAT TTGACTGTTGACAAGTCTTCCACCACAGCCTTCATCTCAACAGCCTGACATCTGACGACTCTGCA

GTITATITCTGTGCAAGAGATGTGACTACGGCCTTGACTTCTGGGGCCAAGGCACCACTCTCACAGTC

V Y F C A R D V T T A L D F W G Q G T T L T V

TCCTCA

FIG. 1B

ANTIBODY	APPARENT K ₁ , M ⁻¹	APPARENT K ₁ , M	
BY ELISA			
D2	5.2 x 10 ⁹	1.9 x 10 ⁻¹⁰	
147	6.5 x 10 ⁹	1.5 x 10 ⁻¹⁰	
K73	9.8 x 10 ⁹	1.0×10^{-10}	
K80	2.3 x 10 ⁹	4.3×10^{-10}	
L102	2.5 x 10 ⁹	4.0×10^{-10}	
L133	1.7 x 10 ⁹	5.9×10^{-10}	
BY BIACore			
H36	3.1 x 10 ¹⁰	3.2 x 10 ⁻¹¹	
143	2.3 x 10 ⁹	4.3 x 10 ⁻¹⁰	
147	3.2 x 10 ⁹	3.1 x 10 ⁻¹⁰	
L133	4.6 x 10 ⁹	2.2 x 10 ⁻¹⁰	
M107	1.1 x 10 ⁹	9.1 x 10 ⁻¹⁰	

FIG. 2

ANTIBODY NAME	% INHIBITION ANTIBODY PREINCUBATED WITH TF/VIIa		
D1	0		
D1B	1		
H31	4		
H36	95		
143	1		
J131	7		
K80	0		
K82	Ō		
K87	1		
L97B	7		
L101	0		
L102	0		
L105	0		
L133	0		
M5	1		
M107	34		

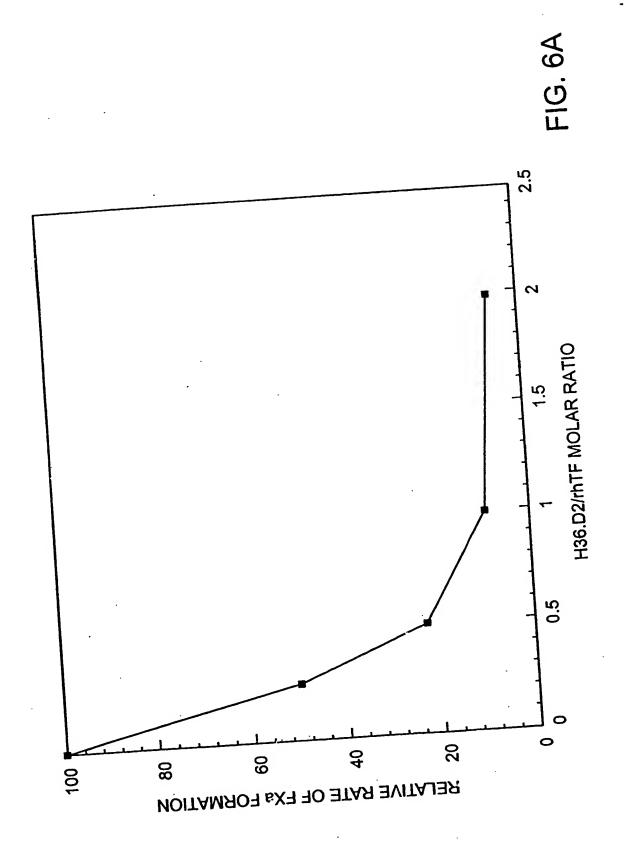
FIG. 3

ANTIBODY NAME	% INHIBITION TF PREINCUBATED WITH ANTIBODY PRIOR TO ADDITION OF VIIa	% INHIBITION TF PREINCUBATED WITH VIIa PRIOR TO ADDITION OF ANTIBODY
D1 D1B H31 H36 I43 J131 K80 K82 K87 L96 L101 L102 L105 L133 M5 M107	15 48 64 0 68 38 12 0 0 0 0 38 14 4 13 0	nd 12.7 21 0 55 11 nd nd nd nd nd nd nd

FIG. 4

[thTF], nM	[H36.D2], nM	H36.D2/mTF MOLAR RATIO	CLOTTING TIME (SECONDS)	% INHIBITION OF thTF FUNCTION
0.0048	0	0	102.3	0
	1.61	335.4	114.3	31.3
	3.23	670.8	121.3	45.8
0.023	0	0	77.6	0
	1.61	70.0	85.3	52.2
	3.23	140.0	91.1	65.2
	6.45	280.4	99.6	73.9
0.092	0	0	49.3	0
	3.23	35.1	65.8	65.2
	6.45	70.1	88.5	90.2
	12.90	140.2	113.3	95.7
0.46	0	0	32.6	0
	6.45	14.0	52.7	82.4
	12.90	28.0	80.2	96.7
	32.30	70.2	117.9	99.3
2.30	0	0	23.9	0
	16.10	7.0	47.1	94.4
	32.30	14.0	95.2	99.7
	64.50	28.0	115.3	99.9
11.52	0	0	22.2	0
	16.10	1.4	30.2	93.4
	32.30	2.8	46.0	98.8
	64.50	5.6	87.6	99.9
	161.30	14.0	114.0	100.0

FIG. 5



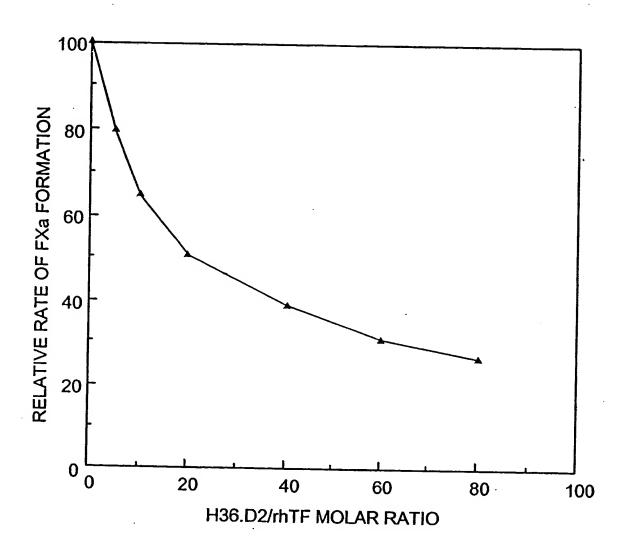


FIG. 6B

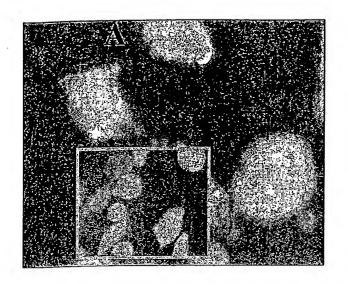
% INHIBITION ID FX AND H36.D2 ARE TED ADDED SIMULTANEOUSLY TO CELLS (TF/FVII)	0	pu	pu	pu	76	78	. 26
% INHIBITION CELLS (TF/FVII) AND H36.D2 PREINCUBATED PRIOR TO FX ADDITION	0	88	92	26	pu	pu	pu
H36.D2 CONCENTRATION (ng)	0	50	100	200	800	1600	3200

FIG. 7

	က <u> </u>	UREA	0	o	0	
	7	UREA	0	Ø	0	
		NATIVE UREA	0	Ø	Ø	
			60ng	30ng	15ng	
	Α	∑⊢			,	
က	8M UREA					
7		8M UREA	0	Ø	0	
-		8M NATIVE UREA	0	Ø	Ø	
		·	2µl, 60ng	1μl, 30Ag	0.5µl, 15ng	

FIG. 8B

FIG. 8A



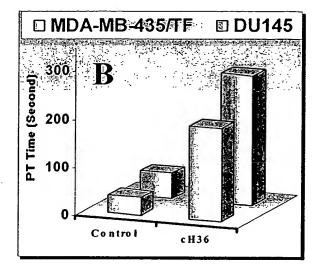


Fig. 9A

Fig. 9B

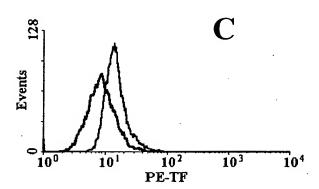


Fig. 9c

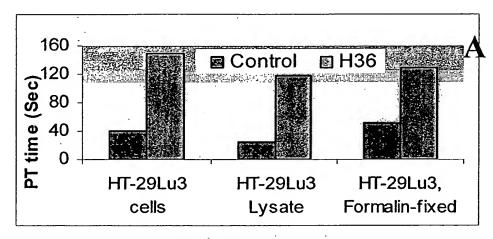


Fig. 10A

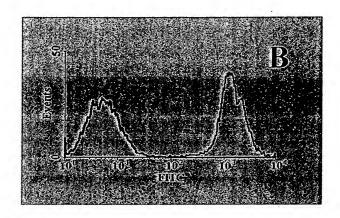


Fig. 10B

Lung Tumor

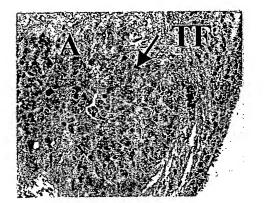


Fig. 11A

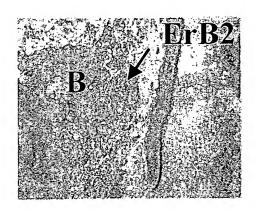


Fig. 11B

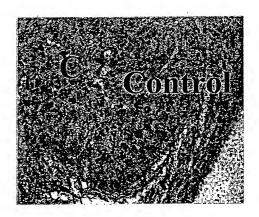


Fig. 11C

Normal Lung

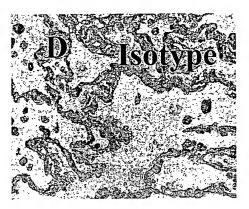


Fig. 11D

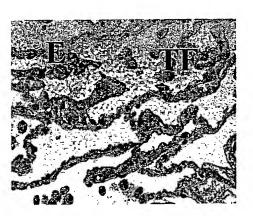
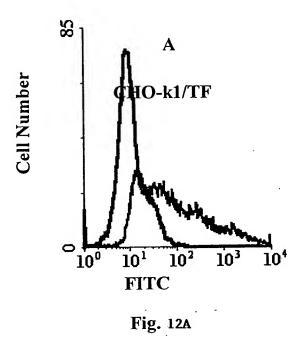
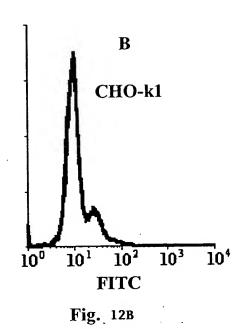


Fig. 11E





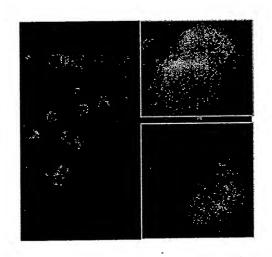


Fig. 12c

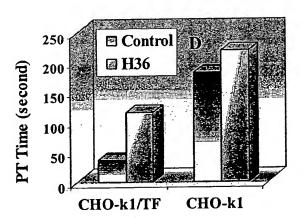


Fig. 12D